



IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

Application of:)
)
D'Amato et al.)
)
 Serial No. **09/899,702**)
)
Filed: July 5, 2001)
)
For: Estrogenic Compounds as)
Antioangiogenic Agents)

DEC 15 2003
 OFFICE OF PETITIONS
 Art Unit: 1616
 Examiner: **Qazi, Sabiha Naim**

SUPPLEMENTAL INFORMATION DISCLOSURE STATEMENT

Commissioner for Patents
 P.O. Box 1450
 Alexandria, VA 22313-1450

Sir:

The citation of information on the attached Form PTO/SB/08 is made pursuant to 37 C.F.R. §§ 1.56, 1.97, and 1.98. A copy of each cited item is enclosed.

As this Information Disclosure Statement is being filed after the issuance of an office action on the merits, pursuant to 37 C.F.R. §1.97(c)(2), the fee of \$180, as set forth in §1.17(p), is enclosed.

The citation of this information does not constitute an admission of priority or that any cited item is available as a reference, or a waiver of any right the applicant may have under applicable statutes, Rules of Practice in patent cases, or otherwise.

Respectfully submitted,

 Robert E. Richards
 Reg. No. 29,105

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 Suite 2800, 1100 Peachtree Street
 Atlanta, Georgia 30309
 (404) 815-6500
 Our Docket: 05213-0910 (43170-260981)

I hereby certify that this correspondence is being deposited with the United States Postal Service as first class mail in an envelope addressed to: Commissioner for Patents, P.O. Box 1450, Alexandria, VA 22313-1450, on 11-26-03.

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Substitute for Form 1449/A/PTO

**INFORMATION DISCLOSURE
STATEMENT BY APPLICANT**

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Complete if Known

Application No.	09/899,702
Filing Date	07/05/2001
First Named Inventor	Robert D'Amato
Group Art Unit	1616
Examiner Name	Qazi, S.
Attorney Docket Number	05213-0910 (43170-260981)

Sheet 1 of 14

U.S. PATENT DOCUMENTS

Examiner Initials	Cite No. ¹	U.S. Patent Document		Name of Patentee or Applicant of Cited Document	Date of Publication of Cited Document MM-DD-YYYY	Pages, Columns, Lines, Where Relevant Passages or Relevant Figures Appear
		Number ⁴	Kind Code ² (if known)			
	1	4,552,758		Murphy et al.	11/12/1985	
	2	4,634,705		DeBernardis et al.	01/06/1987	
	3	4,808,402		Leibovich et al.	02/28/1989	
	4	5,629,340	A	Kuwano et al.	05/13/1977	
	5	5,639,725		O'Reilly et al.	06/17/1997	
	6	5,716,981		Hunter et al.	02/10/1998	
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	8	5,776,704		O'Reilly et al.	07/07/1998	
	9	5,792,845		O'Reilly et al.	08/11/1998	
	10	5,837,682		O'Reilly	11/17/1998	
	11	5,854,205		O'Reilly et al.	12/29/1998	
	12	5,854,221		Cao et al.	12/29/1998	
	13	5,861,372		Folkman et al.	01/19/1999	
	14	5,885,795		O'Reilly et al.	03/23/1999	
	15	6,514,971	B1	Thomas et al.	02/04/2003	

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				Examiner Name	Qazi, S.			
Sheet	2	of	14	Attorney Docket Number	05213-0910 (43170-260981)			
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		Office ³	Number ⁴	Kind Code ⁵ (if known)				
	16	EPO	0166937	A2	Abbott Laboratories	08/01/1986		
	17	DE	27 57 157		Schering AG	12/19/1977		
	18	WO	93/19746	A1	ORION-YH-TYMA OY	10/14/1993		
	19	WO	99/22728	A1	Arch Development Corporation	05/14/1999		
	20	JP	11-209322			01/27/1998		No
	21	JP	58-131978			08/06/1983		No
	22	JP	63-119500			05/24/1988		No

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OTHER PRIOR ART - NON PATENT LITERATURE DOCUMENTS

Examiner Initials	Cite No. ¹	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published	T ²
	23	Author: _____ Title: _____ Publ: Research Plus Catalog Vol/Iss: _____ Pages: 50-58 Date: 1993	
	24	Author: _____ Title: Registry Number: 56933-77-8 Publ: Chemical Abstracts Vol/Iss: _____ Pages: _____ Date: _____	
	25	Author: _____ Title: Registry Number: 56933-78-9 Publ: Chemical Abstracts Vol/Iss: _____ Pages: _____ Date: _____	
	26	Author: _____ Title: Registry Number: 57380-15-1 Publ: Chemical Abstracts Vol/Iss: _____ Pages: _____ Date: _____	
	27	Author: _____ Title: Registry Number: 71782-94-0 Publ: Chemical Abstracts Vol/Iss: _____ Pages: _____ Date: _____	
	28	Author: _____ Title: Registry Number: 71782-95-1 Publ: Chemical Abstracts Vol/Iss: _____ Pages: _____ Date: _____	
	29	Author: _____ Title: Registry Number: 101277-11-6 Publ: Chemical Abstracts Vol/Iss: _____ Pages: _____ Date: _____	

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	30	Author: Title: Registry Number: 101429-40-7 Publ: Chemical Abstracts Vol/Iss: Pages: Date:	RECEIVED DEC 15 2003 OFFICE OF PETITIONS
	31	Author: Title: Registry Number: 162853-20-5 Publ: Chemical Abstracts Vol/Iss: Pages: Date:	
	32	Author: Title: News Article: Hoffman-La Roche Signs \$70 Million Deal with Millenium on Genomics Technology Publ: Genetic Engineering News Vol/Iss: Pages: Date: 4/15/1994	
	33	Author: Title: News Article: Advanced Drug Delivery Systems Peak Interest of Pharmaceutical & Biotech Firms Publ: Genetic Engineering News Vol/Iss: Pages: Date: 4/15/1994	
	34	Author: Title: News Article: Nasal Spray for Treating Bleeding Disorders Publ: Genetic Engineering News Vol/Iss: Pages: Date: 4/15/1994	
	35	Author: ALGIRE, G.H. et al. Title: Vascular reactions of normal and malignant tumors in vivo. I. Vascular reactions of mice to wounds and to normal and neoplastic transplants Publ: Journal of the National Cancer Institute Vol/Iss: 6 Pages: 73-85 Date: 08/1945	
	36	Author: ALIEV et al. Title: 54929q Synthesis of cycloalkyl derivatives of dihydric phenols and their ethers Publ: Chemical Abstracts Vol/Iss: 72 Pages: 370 Date: 1970	

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	37	Author: ANSTEAD et al. Publ: Steroids	Title: The Estradiol Pharmacophore: Ligand Structure-Estrogen Receptor Binding Affinity Relationships and a Model for the Receptor Binding Site	Vol/Iss: 62	Pages: 268-303 Date: 1997
	38	Author: BARNES et al. Publ: Infection and Immunity	Title: Tumor Necrosis Factor Production in Patients with Leprosy	Vol/Iss: 60 (4)	Pages: 1441-1446 Date: 04/1992
	39	Author: BINDRA et al. Publ: Journal of Medicinal Chemistry	Title: Studies in Antifertility Agents. 8. Seco Steroids. 2. 5,6-Secoestradiol and Some Related Compounds	Vol/Iss: 18 (9)	Pages: 921-925 Date: 1975
	40	Author: BOYCE et al. Publ: Unknown	Title: Some Preliminary Synthetical Studies with 5,6,7,8-Tetra-hydro-8-methylindane -1,5-dione	Vol/Iss:	Pages: 4547-4553 Date: 1960
	41	Author: BREM, H. et al. Publ: Journal of Neurosurgery	Title: Interstitial chemotherapy with drug polymer implants for the treatment of recurrent gliomas	Vol/Iss: 74	Pages: 441-446 Date: 03/01/91
	42	Author: BURROWS, N.P. Publ: British Medical Journal	Title: Thalidomide Modifies Disease	Vol/Iss: 307 (6909)	Pages: 939-940 Date: 10/09/93
	43	Author: CAMBIE et al. Publ: J. Chem. Soc.	Title: Aromatic Steroids. Part I. Oxidation Products of 3-Methoxyestra-1,3,5(10)-triene- 17 β -yl Acetate	Vol/Iss:	Pages: 2603-2608 Date: 1968

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	44	Author: COREY et al. Title: Applications of N,N-Dimethylhydrazones to Synthesis. Use in Efficient, Positionally and Stereochemically Selective C-C Bond Formation; Oxidative Hydrolysis to Carbonyl Compounds Publ: <u>Tetrahedron Letters</u> Vol/Iss: 1 Pages: 3-6 Date: 1976				
	45	Author: COREY et al. Title: Facile Conversion of N,N-Dimethylhydrazones to Carbonyl Compounds by Cupric Ion-Catalyzed Hydrolysis Publ: <u>Tetrahedron Letters</u> Vol/Iss: 41 Pages: 36678-3668 Date: 1976				
	46	Author: D'AMATO, R.J. et al. Title: Thalidomide is an Inhibitor of Angiogenesis Publ: <u>Proceedings of the National Academy of Science USA</u> Vol/Iss: 91 Pages: 4082-4085 Date: 04/01/94				
	47	Author: DURANI et al. Title: Seco-Oestradiols and Some Non-Steroidal Oestrogens: Structural Correlates of Oestrogenic Action Publ: <u>Journal of Steroid Biochemistry</u> Vol/Iss: 11 Pages: 67-77 Date: 1979				
	48	Author: EDER et al. Title: Synthese von Ostradiol (in German - No translation available) Publ: <u>Chem. Ber.</u> Vol/Iss: 109 Pages: 2948-2953 Date: 1976				
	49	Author: FEVIG et al. Title: A Short, Stereoselective Route to 16 α -(Substituted-alkyl)estradiol Derivatives Publ: <u>Journal of Organic Chemistry</u> Vol/Iss: 52 Pages: 247-251 Date: 1987				
	50	Author: FIELD et al. Title: Effect of Thalidomide on the Graft versus Host Reaction Publ: <u>Nature</u> Vol/Iss: 211 (5055) Pages: 1308-1310 Date: 09/17/66				

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OTHER PRIOR ART - NON PATENT LITERATURE DOCUMENTS

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	51	Author: FIESER et al. Publ: Organic Synthesis Collective Volume 3 Vol/Iss: 3	Title: N-Methylformanilide Pages: 590-591	Date: DEC 15 2003
	52	Author: FLOHE et al. Publ: Arzneimittel/Forschung (Germany West) Vol/Iss: 31 (2)	Title: Studies on the Hypothetical Relationship of Thalidomide-induced Embryopathy and Collagen Biosynthesis Pages: 315-320	Date: 01/01/81
	53	Author: FOLKMAN et al. Publ: Science Vol/Iss: 221	Title: Angiogenesis Inhibition and Tumor Regression Caused by Heparin or a Heparin Fragment in the Presence of Cortisone Pages: 719-725	Date: 08/19/83
	54	Author: FOLKMAN, J. Publ: New England Journal of Medicine Vol/Iss: 285 (21)	Title: Tumor Angiogenesis: Therapeutic Implications Pages: 1182-1186	Date: 11/18/71
	55	Author: FOLKMAN, J. et al. Publ: Nature Vol/Iss: 339	Title: Induction of Angiogenesis During the Transition from Hyperplasia to Neoplasia Pages: 58-61	Date: 05/04/89
	56	Author: FOLKMAN, J. et al. Publ: Annals of Surgery Vol/Iss: 164(3)	Title: Tumor Behavior in Isolated Perfused Organs in vitro Growth and Metastases of Biopsy Material in Rabbit Thyroid and Canine Intestinal Segment Pages: 491-502	Date: 09/01/66
	57	Author: GASLINI et al. Publ: Journal of Organic Chemistry Vol/Iss: 29 (5)	Title: Reaction of Eugenol with Synthesis Gas. Synthesis of 5,6,7,8-Tetrahydro-3-methoxy-2-naphthol Pages: 1177-1180	Date: 05/1964

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	58	Author: GIMBRONE, M.A. et al. Title: Tumor Growth and Neovascularization: An Experimental Model Using the Rabbit Cornea Publ: <u>Journal of the National Cancer Institute</u> Vol/Iss: 52(2) Pages: 413-427 Date: 02/74	
	59	Author: GIMBRONE, M.A. et al. Title: Tumor dormancy <i>in vivo</i> by Prevention of Neovascularization Publ: <u>Journal of Experimental Medicine</u> Vol/Iss: 136 Pages: 261-276 Date: 1972	
	60	Author: GONZALEZ et al. Title: Synthesis and Pharmacological Evaluation of 8 α Estradiol Derivatives Publ: <u>Steroids</u> Vol/Iss: 40 (2) Pages: 171-187 Date: 9/1982	
	61	Author: GROSS, J.L. et al. Title: Modulation of Solid Tumor Growth <i>in vivo</i> by bFGF (Abstract only) Publ: <u>Proceedings of the American Association of Cancer Research</u> Vol/Iss: 31 Pages: 79 Date: 03/90	
	62	Author: GUNZLER, V. Title: Thalidomide-A Therapy for the Immunological Consequences of HIV Infection? Publ: <u>Medical Hypothesis</u> Vol/Iss: 30 (2) Pages: 105-109 Date: 10/89	
	63	Author: GUPTA et al. Title: Antifertility Agents. XIV. Secosteroids. VII. Synthesis of 2 α and 2 β , 6 β -dimethyl- 3 β -(p-hydroxyphenyl)-trans-bicyclo[4.3.0]nonan-7-ones and some related compounds (Abstract only) Publ: <u>Indian Journal of Chemistry</u> Vol/Iss: 13 (7) Pages: 759-760 Date: 1975	
	64	Author: GUPTA et al. Title: Studies in Antifertility Agents. Part XVIII. 2 α 6 β -Diethyl-3 β -(p-hydroxyphenyl)-trans-bicyclo[4.3.0]nonan-7 β -ol and 6 β -methyl-3 β -(p-hydroxyphenyl)-2 α -propyl- trans-bicyclo[4.3.0]nonan-7 β -ol (Abstract only) Publ: <u>Indian Journal of Chemistry</u> Vol/Iss: 19B (10) Pages: 886-890 Date: 1980	

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	65	Author: GUTIERREZ-RODRIGUEZ et al. Title: Treatment of Refractory Rheumatoid Arthritis - The Thalidomide Experience Publ: The Journal of Rheumatology Vol/Iss: 16 (2) Pages: 158-163 Date: 02/89	
	66	Author: GUTIERREZ-RODRIGUEZ, O. Title: Thalidomide - A Promising New Treatment for Rheumatoid Arthritis Publ: Arthritis and Rheumatism Vol/Iss: 27 (10) Pages: 1118-1121 Date: 10/84	
	67	Author: HANDLEY et al. Title: Chronic bullous disease of childhood and ulcerative colitis Publ: British Journal of Dermatology Vol/Iss: 127 (40) Pages: 67-68 Date: 07/01/92	
	68	Author: HENEY et al. Title: Thalidomide treatment for chronic graft-versus-host disease Publ: British Journal of Haematology Vol/Iss: 78 (1) Pages: 23-27 Date: 05/91	
	69	Author: HORI, A. et al. Title: Suppression of Solid Tumor Growth by Immunoneutralizing Monoclonal Antibody Against Human Basic Fibroblasts Growth Factor Publ: Cancer Research Vol/Iss: 51 Pages: 6180-6184 Date: 11/15/91	
	70	Author: IMAMURA et al. Title: Method for Manufacture of Dihydric Phenols (Abstract only) Publ: USPATFULL 76:20259 US 3,950,437 Vol/Iss: Pages: Date: 04/13/1976	
	71	Author: INGBER, D. et al. Title: Synthetic analogues of fumagillin that inhibit angiogenesis and suppress tumor growth Publ: Nature Vol/Iss: 348 Pages: 555-557 Date: 12/06/90	

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	72	Author: JHINGRAN et al. Title: Studies in Antifertility Agents - Part XLI: Secosteroids-x: Syntheses of Various Stereoisomers of (+)-2,6 β -diethyl-7 α -ethynyl-3-(p-hydroxyphenyl)-trans-bicyclo [4.3.0]nonan-7 β -ol. Publ: <u>Steroids</u> Vol/Iss: 42 (6) Pages: 627-634 Date: 1983	
	73	Author: KARWAT Title: Separation and Recovery of Hydrogen Sulfide from Hydrocarbon Mixture Publ: <u>Caplus DE 1103310</u> Vol/Iss: Pages: Date: 09/02/1959	
	74	Author: KIM, K.J. et al. Title: Inhibition of Vascular Endothelial Growth Factor-induced Angiogenesis Suppresses Tumor Growth In Vivo Publ: <u>Nature</u> Vol/Iss: 362 Pages: 841-844 Date: 04/29/93	
	75	Author: KNIGHTON, D. et al. Title: Avascular and Vascular Phases of Tumour Growth in the Chick Embryo Publ: <u>British Journal of Cancer</u> Vol/Iss: 35 Pages: 347-356 Date: 1977	
	76	Author: KOLE et al. Title: Studies in Antifertility Agents. 11. Secosteroids.5.Synthesis of 9,11-Secoestradiol Publ: <u>Journal of Medicinal Chemistry</u> Vol/Iss: 18 (7) Pages: 765-766 Date: 1975	
	77	Author: LEWIS, Richard J. Title: Publ: <u>Hawley's Condensed Chemical Dictionary</u> Vol/Iss: Pages: 577 Date: 01/1993	
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¹Unique citation designation number. ²See attached Kinds of U.S. Patent Documents. ³Enter Office that issued the document, by the two-letter code (WIPO Standard ST.3). ⁴For Japanese patent document, the indication of the year of the reign of the Emperor must precede the serial number of the patent document. ⁵Kind of document by the appropriate symbols as indicated on the document under WIPO Standard ST. 16 if possible. ⁶Applicant is to place a check mark here if English language translation is attached.

Substitute for Form 1449/A/PTO				Complete if Known	
INFORMATION DISCLOSURE STATEMENT BY APPLICANT (use as many sheets as necessary)				Application No.	09/899,702
				Filing Date	07/05/2001
				First Named Inventor	Robert D'Amato
				Group Art Unit	1616
				Examiner Name	Qazi, S.
Sheet	11	of	14	Attorney Docket Number	05213-0910 (43170-260981)

OTHER PRIOR ART - NON PATENT LITERATURE DOCUMENTS

Examiner Initials	Cite No. ¹	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published	T ²
	79	Author: LIEN, W. et al. Title: The blood supply of experimental liver metastases. II. A Microcirculatory study of the normal and tumor vessels of the liver with the use of perfused silicone rubber Publ: <u>Surgery</u> Vol/Iss: 68 (2) Pages: 334-340 Date: 08/1970	
	80	Author: LIU et al. Title: Total Synthesis of (+)- $\Delta^{9(12)}$ Capnellene Publ: <u>Tetrahedron Letters</u> Vol/Iss: 26 (40) Pages: 4847-4850 Date: 1985	
	81	Author: MARO et al. Title: Mechanism of Polar Body Formation in the Mouse Oocyte: An Interaction Between the Chromosomes, the Cytoskeleton and the Plasma Membrane Publ: <u>Journal of Embryology and Experimental Morphology</u> Vol/Iss: 92 Pages: 11-32 Date: 1986	
	82	Author: MEZA et al. Title: Managing the Gastrointestinal Complications of AIDS Publ: <u>Drug Therapy</u> Vol/Iss: 23 (11) Pages: 74-83 Date: 11/1993	
	83	Author: NAAFS et al. Title: Thalidomide Therapy An Open Trial Publ: <u>International Journal of Dermatology</u> Vol/Iss: 24 (2) Pages: 131-134 Date: 03/1985	
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	86	Author: NGUYEN, M. et al. Title: Elevated Levels of the Angiogenic Peptide Basic Fibroblast Growth Factor in Urine of Bladder Cancer Patients Publ: <u>Journal of the National Cancer Institute</u> Vol/Iss: 85 (3) Pages: 241-242 Date: 02/03/93	
	87	Author: POWELL et al. Title: Investigation and Treatment of Orogenital Ulceration; studies on a Possible Mode of Action of Thalidomide Publ: <u>British Journal of Dermatology</u> Vol/Iss: 113 Supp. 28 Pages: 141-144 Date: 07/1985	
	88	Author: ROMANELLI et al. Title: Ethyl-p-Dimethylaminophenylacetate Publ: <u>Organic Synthesis</u> Vol/Iss: 5 Pages: 552 Date:	
	89	Author: SEEGERs et al. Title: The Mammalian Metabolite, 2-methoxyestradiol, Affects P53 Levels and Apoptosis Induction in Transformed Cells but Not in Normal Cells (Abstract only) Publ: <u>Journal of Steroid Biochemistry and Molecular Biology</u> Vol/Iss: 62 (4) Pages: 253-267 Date: 07/1997	
	90	Author: SHAH et al. Title: (+/-)-(N-alkylamino)benzazepine Analogs: Novel Dopamine D1 Receptor Antagonists Publ: <u>Journal of Medicinal Chemistry</u> Vol/Iss: 38 (21) Pages: 4284-4293 Date: 10/13/95	
	91	Author: SIDKY et al. Title: Inhibition of Angiogenesis by Interferons: Effects on Tumor- and Lymphocyte-induced Vascular Responses Publ: <u>Cancer Research</u> Vol/Iss: 47 Pages: 5155-5161 Date: 10/01/87	
	92	Author: SRIVASTAVA, A. et al. Title: The Prognostic Significance of Tumor Vascularity in Intermediate-Thickness (0.76-4.0 mm Thick) Skin Melanoma Publ: <u>American Journal of Pathology</u> Vol/Iss: 133 (2) Pages: 419-424 Date: 11/1988	

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	93	Author: STARKOV et al. Title: Mono- and Dialkylaton of Guaiacol by Olefins on KU-2 Cation Exchanger (Abstract only) Publ: Zhurnal Prikladnoi Khimii Vol/Iss: 41 (3) Pages: 688-690 Date: 1968	
	94	Author: TAYLOR, S. et al. Title: Protamine is an Inhibitor of Angiogenesis Publ: Nature Vol/Iss: 297 Pages: 307-312 Date: 05/27/82	
	95	Author: TREMBLAY et al. Title: A Convenient Synthetic Method for Alpha-Alkylation of Steroidal 17-Ketone: Preparation of 16 β -(THPO-Heptyl)-Estradiol Publ: Synthetic Communications Vol/Iss: 25 (16) Pages: 2483-2495 Date: 1995	
	96	Author: TREMBLAY et al. Title: Synthesis of 16-(Bromoalkyl)-Estradiols Having Inhibitory Effect on Human Placental Estradiol 17 β -Hydroxysteroid Dehydrogenase (17 β HSD Type 1) Publ: Bioorganic & Medicinal Chemistry Vol/Iss: 3 (5) Pages: 505-523 Date: 1995	
	97	Author: VICENTE et al. Title: In Vitro Activity of Thalidomide Against Mycobacterium avium Complex Publ: Archives of Internal Medicine Vol/Iss: 153 (4) Pages: 534 Date: 02/22/93	
	98	Author: WEIDNER, N. et al. Title: Tumor angiogenesis: A New Significant and Independent Prognostic Indicator in Early-Stage Breast Carcinoma Publ: Journal of the National Cancer Institute Vol/Iss: 84 Pages: 1875-1887 Date: 12/16/92	
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	100	Author: WEIDNER, N. et al. Title: Tumor Angiogenesis and Metastasis-Correlation in Invasive Breast Carcinoma Publ: <u>New England Journal of Medicine</u> Vol/Iss: 324 (1) Pages: 1-8 Date: 01/03/91	
	101	Author: WELSCH et al. Title: Staphylostatic Activity of Some New Diphenols, Napthols, and Chalcones (Abstract only) Publ: <u>Experientia</u> Vol/Iss: 11 Pages: 350-351 Date: 1955	
	102	Author: WHITE et al. Title: Treatment of Pulmonary Hemangiomas with Recombinant Interferon Alfa-2a Publ: <u>The New England Journal of Medicine</u> Vol/Iss: 32 (18) Pages: 1197-1200 Date: 05/04/89	

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